

What is claimed is:

- 1           1. A method of analyzing tissue, the method comprising:  
2           illuminating a tissue with coherent light;  
3           receiving light reflected from the tissue at a detector to form a series of speckle  
4           patterns; and  
5           analyzing changes in the speckle patterns at time intervals sufficient to measure  
6           changes caused by Brownian motion of objects within the tissue.
- 1           2. The method of claim 1, further comprising compensating for extrinsic motion to  
2           isolate the Brownian motion.
- 1           3. The method of claim 1, wherein the tissue is *in vivo*.
- 1           4. The method of claim 1, wherein the tissue is internal tissue.
- 1           5. The method of claim 4, wherein the illuminating step comprises providing an  
2           invasive device coupled to a light source, passing the device into a patient, placing the device  
3           in proximity to the tissue, and shining coherent light from the light source onto the tissue.
- 1           6. The method of claim 5, wherein the invasive device is selected from the group  
2           consisting of a catheter, an endoscope, and a laparoscope.
- 1           7. The method of claim 5, wherein the placing step includes placing the device in  
2           direct contact with the tissue.
- 1           8. The method of claim 5, wherein the invasive device comprises a catheter having a  
2           first fiber that transmits light from the light source to the tissue, and a fiber array that receives  
3           light remitted from the tissue.
- 1           9. The method of claim 1, wherein the coherent light comprises laser light.

1           10. The method of claim 1, wherein the speckle pattern is a far field image formed at  
2           the detector.

1           11. The method of claim 1, wherein the analyzing step comprises comparing each of  
2           the series of speckle patterns to a reference speckle pattern, and quantifying the differences  
3           between each pattern and the reference pattern.

1           12. The method of claim 11, wherein the analyzing step comprises digitizing each of  
2           the speckle patterns, and the quantifying step comprises evaluating a maximum cross-  
3           correlation between each pattern and the reference pattern.

1           13. The method of claim 12, wherein the analyzing step further comprises  
2           determining a decorrelation rate for the speckle patterns.

1           14. The method of claim 1, wherein the analyzing step further comprises analyzing  
2           spatial characteristics of the speckle pattern to deduce structural characteristics of the tissue.

1           15. The method of claim 14, wherein the illuminating step comprises illuminating  
2           multiple sections of the tissue in succession, the receiving step comprises forming a separate  
3           series of speckle patterns for each respective section of the tissue, and the analyzing step  
4           comprises analyzing each separate series of speckle patterns and comparing the separate  
5           series to deduce structural differences between the respective sections of the tissue.

1           16. The method of claim 2, wherein compensating for extrinsic motion comprises  
2           performing the receiving step during a diastole of a heartbeat.

1           17. The method of claim 2, wherein the receiving step comprises gathering reflected  
2           light at a light receptor and transmitting the gathered light to the detector, and wherein  
3           compensating for extrinsic motion includes coupling the receptor to the tissue.

1           18. The method of claim 2, wherein compensating for extrinsic motion includes  
2           excluding changes in the speckle patterns caused by non-random motion during the analysis  
3           step.

1           19. The method of claim 2, wherein extrinsic motion results from blood flow  
2           between the tissue and the reflector, and the compensating step comprises replacing the blood  
3           with a transparent solution.

1           20. The method of claim 1, wherein the tissue comprises atherosclerotic plaque.

1           21. A method of determining the susceptibility to rupture of an atherosclerotic plaque  
2           having a lipid pool and a fibrous cap, the method comprising:  
3           illuminating the plaque with coherent light;  
4           receiving light reflected from the plaque at a detector to form a series of speckle  
5           patterns;  
6           gathering speckle pattern data at time intervals sufficient to measure Brownian  
7           motion within the lipid pool; and  
8           assessing the plaque's vulnerability to rupture from the amount of Brownian motion.

1           22. The method of claim 21, further comprising analyzing spatial characteristics of  
2           the speckle pattern data to determine structural characteristics of the plaque.

1           23. The method of claim 22, wherein the analyzing step comprises assessing the  
2           thickness of the fibrous cap.

1           24. The method of claim 23, wherein a plaque is considered vulnerable to rupture if  
2           the thickness of the fibrous cap is less than about 60 microns.

1           25. The method of claim 22, wherein the analyzing step comprises assessing the  
2           viscosity of the lipid pool.

1           26. The method of claim 25, wherein the plaque is considered vulnerable to rupture if  
2           the viscosity of the lipid pool has a time constant of less than about 200 milliseconds.

1           27. The method of claim 25, wherein the plaque is considered likely to rupture if the  
2           viscosity of the lipid pool has a time constant of less than about 100 milliseconds.

1           28. A method of detecting a vulnerable atherosclerotic plaque having a lipid pool and  
2           a fibrous cap within a blood vessel, the method comprising:  
3           illuminating a segment of the blood vessel *in vivo* with coherent light;  
4           receiving light reflected from the interior vessel wall of the segment at a detector to  
5           form a series of speckle patterns;  
6           gathering speckle pattern data at time intervals sufficient to measure Brownian  
7           motion within the interior vessel wall; and  
8           comparing the speckle pattern data to a known speckle pattern for a normal blood  
9           vessel and a known speckle pattern for an atherosclerotic plaque;  
10          wherein speckle pattern data corresponding to a speckle pattern for an atherosclerotic  
11          plaque indicates the segment of the blood vessel contains an atherosclerotic plaque.

1           29. The method of claim 28, further comprising analyzing spatial characteristics of  
2           the speckle pattern data to determine structural characteristics of the plaque.

1           30. The method of claim 29, wherein the analyzing step comprises assessing the  
2           thickness of the fibrous cap.

1           31. The method of claim 30, wherein a plaque is considered vulnerable to rupture if  
2           the thickness of the fibrous cap is less than about 60 microns.

1           32. The method of claim 29, wherein the analyzing step comprises assessing the  
2           viscosity of the lipid pool.

1           33. The method of claim 32, wherein the plaque is considered vulnerable to rupture if  
2     the viscosity of the lipid pool has a time constant of less than about 200 milliseconds.

1           34. The method of claim 32, wherein the plaque is considered likely to rupture if the  
2     viscosity of the lipid pool has a time constant of less than about 100 milliseconds.

1           35. A fiber optic probe for detecting speckle patterns in a sample, the probe  
2     comprising  
3           a catheter including a rotatable inner shaft and a transparent outer sheath;  
4           a fiber array housed within the shaft and comprising a central optical fiber for  
5     transmitting incident light to the sample and multiple optical fibers for transmitting light  
6     remitted from the sample; and  
7           a mirror arranged near a distal end of the shaft to reflect light passing through the  
8     fiber array onto a sample outside the transparent outer sheath and back from the sample  
9     through the fiber array.

1           36. The fiber optic probe of claim 35, wherein the shaft can rotate 360 degrees within  
2     the sheath.

1           37. The fiber optic probe of claim 35, further comprising an inflatable balloon  
2     connected to the sheath.

1           38. An optical system for detecting speckle patterns in a sample, the system  
2     comprising  
3           a fiber optic probe of claim 35;  
4           a coherent light source connected to the central optical fiber within the fiber array;  
5           a detector to receive light remitted from the sample; and  
6           a processor to process the remitted light and to analyze speckle patterns remitted from  
7     the sample.

1           39. The system of claim 38, wherein the processor comprises a reference speckle  
2 pattern.

1           40. The system of claim 38, wherein the processor comprises an analog-digital  
2 converter to convert the analog remitted light into a digital signal.